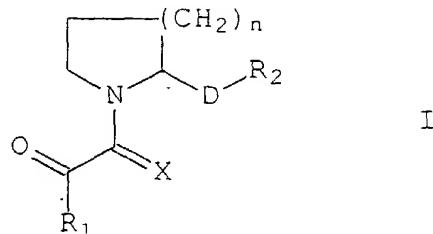


What is claimed is:

1. A compound having the formula (I):



5 where

n is 1-3;

X is either O or S;

10 R₁ is selected from the group consisting of C₁-C₉, straight or branched chain alkyl, C₂-C₉, straight or branched chain alkenyl, aryl, heteroaryl, carbocycle, or heterocycle;

D is a bond, or a C₁-C₁₀ straight or branched chain alkyl, C₂-C₁₀ alkenyl or C₂-C₁₀ alkynyl;

15 R₂ is a carboxylic acid or a carboxylic acid isostere; and wherein said alkyl, alkenyl, alkynyl, aryl, heteroaryl, carbocycle, heterocycle, or carboxylic acid isostere is optionally substituted with one or more substituents selected from R³ and Z, where

20 R³ and Z are independently hydrogen, hydroxy, halo, haloalkyl, thiocarbonyl, alkoxy, alkenoxy, alkylaryloxy, aryloxy, arylalkyloxy, cyano, nitro, imino, alkylamino, aminoalkyl, sulfhydryl, thioalkyl, alkylthio, sulfonyl, C₁-C₆ straight or branched chain alkyl, C₂-C₆ straight or branched chain alkenyl or alkynyl, aryl, aralkyl, heteroaryl, carbocycle, heterocycle, or CO₂R⁷ where R⁷ is hydrogen or C₁-C₉, straight or branched chain alkyl or C₂-C₉, straight or branched chain alkenyl;

25 or a pharmaceutically acceptable salt, ester, or solvate

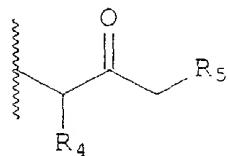
thereof;

provided that:

when n=1, and D is a bond, and R₂ is COOH,
then R₁ is not C₁-C₉ straight or branched chain alkyl, C₂-C₉
straight or branched chain alkenyl, C₅-C, cycloalkyl, C₅-C,
5 cycloalkenyl, phenylamine, 2-(3,4-dichlorophenyl)ethyl,
hydroxy, ethoxy, benzyl, or Ar₁, where Ar₁ is 1-naphthyl, 2-
naphthyl, 2-indolyl, 3-indolyl, 2-furyl, 3-furyl, 2-
thiazolyl, 2-thienyl, 3-thienyl, 1-pyridyl, 2-pyridyl, 3-
10 pyridyl, 4-pyridyl, or phenyl, and wherein said alkyl,
alkenyl, cycloalkyl, cycloalkenyl, or Ar₁ are optionally
substituted with one or more substituents selected from the
group consisting of hydrogen, halo, hydroxyl, nitro,
trifluoromethyl, C₁-C₉ straight or branched alkyl, C₂-C₉,
straight or branched alkenyl, C₁-C₄ alkoxy, C₂-C₄ alkenyloxy,
phenoxy, benzyloxy, COOH, and amino;

further provided that:

when n=1, and D is the carboxylic acid
isostere -CONZ(R³), and Z is hydrogen or C₁-C₆ alkyl, and
R³ is phenyl, or C₂-C₆ straight or branched chain alkyl or
alkenyl, wherein said alkyl is unsubstituted or substituted
in one or more positions with Ar₂ as defined below, C₃-C₈
cycloalkyl, cycloalkyl connected by methyl or a C₂-C₆
straight or branched chain alkyl or alkenyl chain, C₁-C₄
25 alkyl ester, or Ar₃, where Ar₃ is selected from the group
consisting of 2-indolyl, 3-indolyl, 2-furyl, 3-furyl, 2-
thiazolyl, 2-thienyl, 3-thienyl, 2-pyridyl, 3-pyridyl, 4-
pyridyl, or phenyl, having one to three substituents
30 independently selected from the group consisting of hydrogen,
halo, hydroxy, nitro, trifluoromethyl, C₁-C₆ straight or
branched alkyl, C₂-C₆ straight or branched alkenyl, C₁-C₄
alkoxy, C₂-C₄ alkenyloxy, phenoxy, benzyloxy, and amino;
wherein said alkyl ester is optionally substituted with
phenyl; or R³ is the fragment:



where R_4 is selected from the group consisting of straight or branched chain C_1-C_8 alkyl optionally substituted with C_3-C_8 cycloalkyl, benzyl, or Ar_2 , as defined below, and where R_2 is $COOZ$ or $CONR^6$, where R^6 is selected from the group consisting of hydrogen, C_1-C_6 straight or branched alkyl, and C_2-C_6 straight or branched alkenyl, and where R_5 is selected from the group consisting of phenyl, benzyl, C_1-C_6 straight or branched alkyl, and C_2-C_6 straight or branched alkenyl, where said alkyl or alkenyl is optionally substituted with phenyl; then R_1 is not C_1-C_8 straight or branched chain alkyl, C_2-C_9 straight or branched chain alkenyl, substituted thiophene, or C_1-C_4 alkoxy, wherein said alkyl or alkenyl is optionally substituted in one or more positions with C_3-C_8 cycloalkyl, C_5-C_7 cycloalkenyl, or Ar_2 , where Ar_2 is defined below, where said alkyl, alkenyl, cycloalkyl or cycloalkenyl groups may be optionally substituted with C_1-C_4 alkyl, C_1-C_4 alkenyl, or hydroxy, and where Ar_2 is 1-naphthyl, 2-naphthyl, 2-indolyl, 3-indolyl, 2-furyl, 3-furyl, 2-thienyl, 3-thienyl, 2-pyridyl, 3-pyridyl, 4-pyridyl, or phenyl, having one to three substituents selected from the group consisting of hydrogen, halo, hydroxy, nitro, trifluoromethyl, C_1-C_6 straight or branched alkyl, C_2-C_6 straight or branched alkenyl, C_1-C_4 alkoxy, C_2-C_4 alkenyloxy, phenoxy, benzyloxy, and amino; further provided that:
when $n=1$, and X is O , and D is a bond, and R_2 is $-CONH_2$, then R_1 is not methyl, ethyl, iso-propyl, iso-butyl, iso-pentyl, 4-methylpentyl, indolyl, phenyl, or hydroxyphenyl;
further provided that:

when n=1, and X is O, and D is a bond, and R₂ is cyano, then R₁ is not methyl;

further provided that:

when n=2, and X is O, and D is a bond, and R₂ is CONZ(R³), and R₁ is ethoxy, then R³ or Z is not halo-substituted phenyl;

further provided that:

when n=2, and X is O, and D is a bond, and R₂ is CONZ(R³) and R₁ is substituted thiophene or tetrahydropyranoxy, or methoxy, then R³ or Z is not C₁-C₄ alkyl ester substituted ethyl;

further provided that:

when n=2, and X is O, and D is a bond, and R₂ is CONZ(R³) and R₁ is ethoxy, then R³ or Z is not 4-chlorophenyl;

further provided that:

when n=2, and X is O, and D is a bond, and R₂ is CONZ(R³) and R₁ is cyclohexyl, then R³ or Z is not ethyl or propyl substituted with phenyl;

further provided that:

when D is CH₂, then R₂ is not -OMe, -NHMe, or substituted -NHcyclohexyl;

further provided that:

when D is CH₂, and R₂ is -OH, then R₁ is not phenyl or pyrrolidinemethanol;

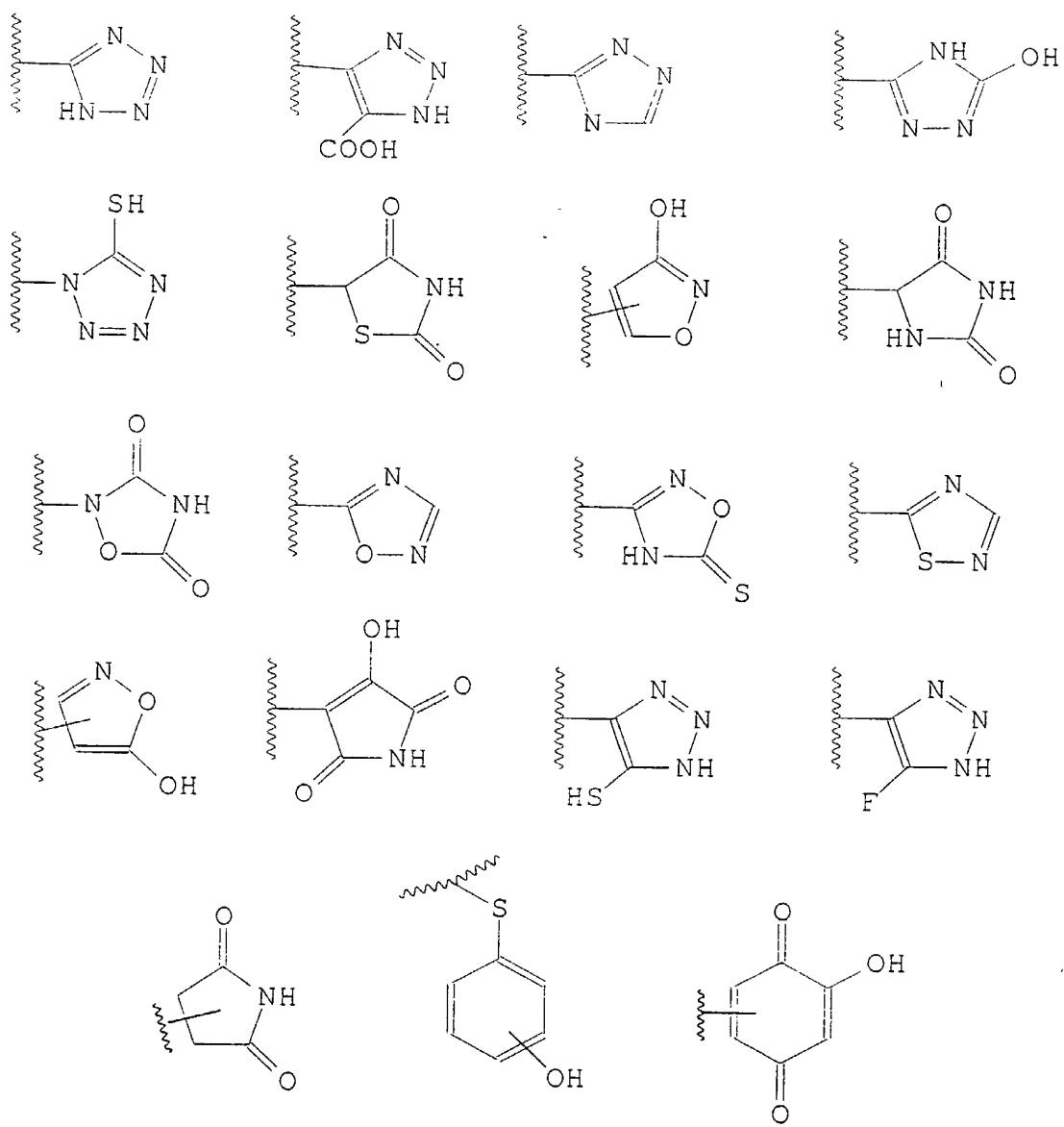
further provided that:

when n=2, and X is O, and D is a bond, and R₂ is COOH, then R₁ is not methyl, tert-butyl, 1,1-dimethyl-2-methyl-propyl, 1,1-dimethyl-propyl, methoxy, ethoxy, phenyl, tetrahydropyranoxy substituted C₄-C₆ alkyl, 1-methyl-1-methoxyamide, 1-methylcyclohexyl, 3-iodophenyl, 3-methyl ester-cyclopentyl, 1,1-dimethyl-6-phenyl-hex-3,5-dioxy, or trimethoxyphenyl.

2. The compound of claim 1, wherein R₂ is a carbocycle or heterocycle containing any combination of CH₂, O, S, or N in

any chemically stable oxidation state, wherein any of the atoms of said ring structure are optionally substituted in one or more positions with R³.

5 3. The compound of claim 1, wherein R₂ is selected from the group consisting of:



where the atoms of said ring structure may be optionally substituted at one or more positions with R³.

4. The compound of claim 1, wherein R₂ is selected from the
5 group consisting of:

-COOH; -SO₃H; -SO₂HNR³; -PO₂(R³)₂; -CN; -PO₃(R³)₂; -OR³; -SR³;
-NHCOR³; -N(R³)₂; -CONZ(R³); -CONH(O)R³; -CONHNHSO₂R³;
-COHNSO₂R³; and -CONR³CN.

10 5. The compounds, (2S)-1-(1,2-dioxo-3,3-dimethylpentyl)-2-hydroxymethylpyrrolidine; (2S)-1-(1,2-dioxo-3,3-dimethylpentyl)-2-pyrrolidinetetrazole; (2S)-1-(1,2-dioxo-3,3-dimethylpentyl)-2-pyrrolidinecarbonitrile; and (2S)-1-(1,2-dioxo-3,3-dimethylpentyl)-2-aminocarbonyl piperidine; and
15 compounds 1-25, 27, 28, 31-33, and 35-136 of Tables I, II, and III.

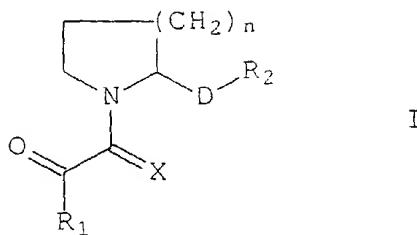
6. The compound 1-[2-[3-(4-Fluorophenyl)(1,2,4-oxadiazol-5-yl)]pyrrolidinyl]-3,3-di-methylpentane-1,2-dione.

20 7. The compound 3,3-Dimethyl-1-[2-(3-methyl(1,2,4-oxadiazol-5-yl))pyrrolidinyl]pentane-1,2-dione.

8. A pharmaceutical composition, comprising:

25 a) an effective amount of an N-heterocyclic carboxylic acid or carboxylic acid isostere; and
b) a pharmaceutically acceptable carrier.

30 9. The pharmaceutical composition of claim 8, wherein the N-heterocyclic carboxylic acid or carboxylic acid isostere comprises a compound of formula (I):



where

n is 1-3;

-- X is either O or S;

5 R₁ is selected from the group consisting of C₁-C₉, straight or branched chain alkyl or alkenyl, C₂-C₉, straight or branched chain alkenyl, aryl, heteroaryl, carbocycle, or heterocycle;

D is a bond, or a C₁-C₁₀ straight or branched chain alkyl, 10 C₂-C₁₀ alkenyl or C₁-C₁₀ alkynyl;

R₂ is carboxylic acid or a carboxylic acid isostere; and

wherein said alkyl, alkenyl, alkynyl, aryl, heteroaryl, carbocycle, or heterocycle is optionally substituted with one 15 or more substituents selected from R³, where

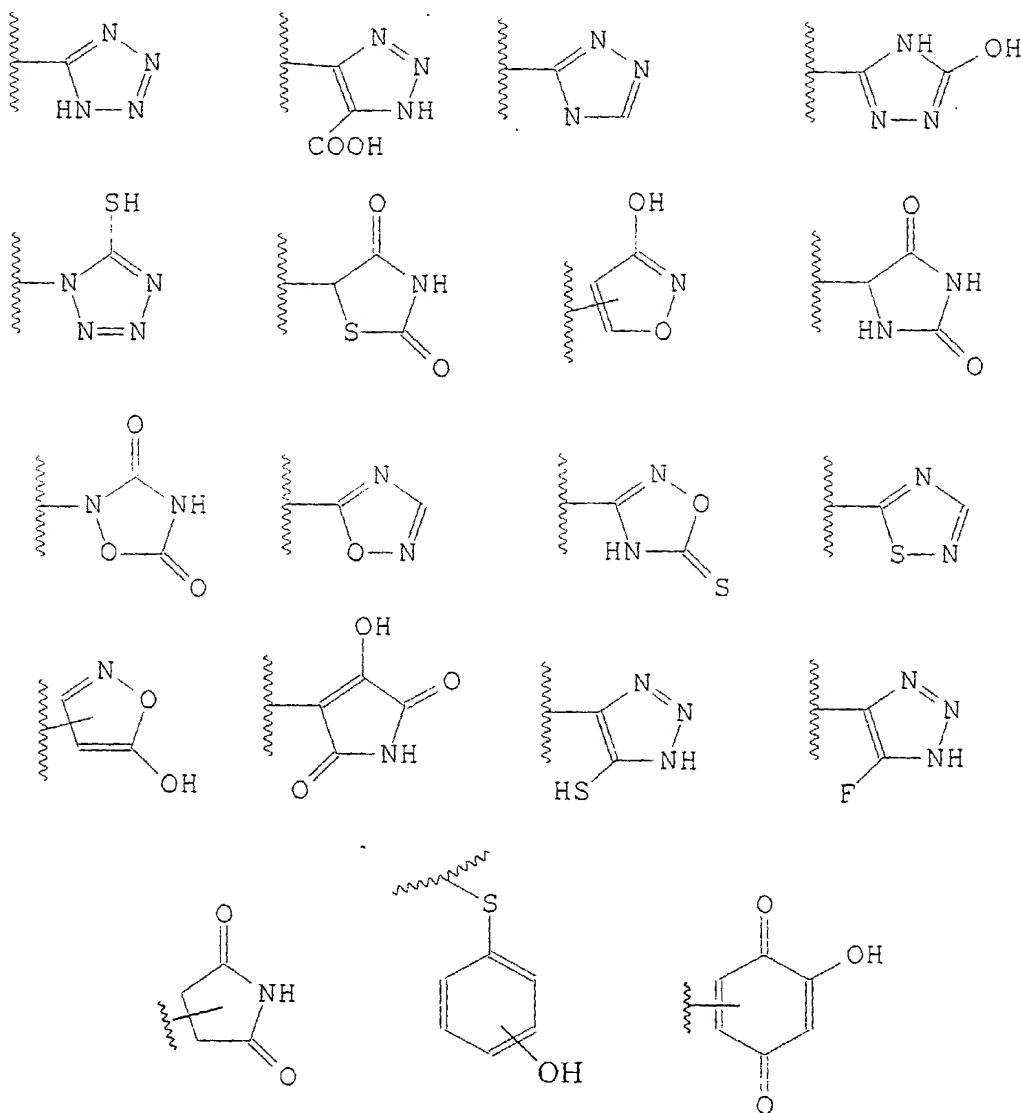
R³ is hydrogen, hydroxy, halo, haloalkyl, thiocarbonyl, alkoxy, alkenoxy, alkylaryloxy, aryloxy, arylalkyloxy, cyano, nitro, imino, alkylamino, aminoalkyl, sulfhydryl, thioalkyl, alkylthio, sulfonyl, C₁-C₆ straight or branched chain alkyl, 20 C₂-C₆ straight or branched chain alkenyl or alkynyl, aryl, aralkyl, heteroaryl, carbocycle, heterocycle, and CO₂R⁷ where R⁷ is hydrogen or C₁-C₉, straight or branched chain alkyl or C₂-C₉, straight or branched chain alkenyl; or a pharmaceutically acceptable salt, ester, or solvate thereof.

10. The pharmaceutical composition of claim 9, wherein R₂ is

a carbocycle or heterocycle containing any combination of CH₂, O, S, or N in any chemically stable oxidation state, wherein any of the atoms of said ring structure are optionally substituted in one or more positions with R³.

5

11. The pharmaceutical composition of claim 9, wherein R₂ is selected from the following group:



where the atoms of said ring structure may be optionally substituted at one or more positions with R³.

12. The pharmaceutical composition of claim 9, wherein R₂ is
5 selected from the group consisting of:
-COOH; -SO₃H; -SO₂HNR³; -PO₂(R³)₂; -CN; -PO₃(R³)₂; -OR³; -SR³;
-NHCOR³; -N(R³)₂; -CON(R³)₂; -CONH(O)R³; -CONHNHSO₂R³;
-COHNSO₂R³; and -CONR³CN..

10 13. The pharmaceutical composition of claim 9, wherein the
N-heterocyclic carboxylic acid or carboxylic acid isostere
compound is selected from the group consisting of compounds
1-139.

15 14. The pharmaceutical composition of claim 8, further
comprising a neurotrophic factor different from formula (I).

15. The pharmaceutical composition of claim 14, wherein said
neurotrophic factor different from formula (I) is selected
20 from neurotrophic growth factor, brain derived growth factor,
glial derived growth factor, ciliary neurotrophic factor,
insulin growth factor and active truncated derivatives
thereof, acidic fibroblast growth factor, basic fibroblast
growth factor, platelet-derived growth factors, neurotropin-3
25 and neurotropin 4/5.

16. A method of treating a neurological disorder in an
animal, comprising:
30 administering to the animal an effective amount of an
N-heterocyclic carboxylic acid or carboxylic acid isostere
to stimulate growth of damaged peripheral nerves or to
promote neuronal regeneration.

35 17. The method of claim 16, wherein the neurological
disorder is selected from the group consisting of peripheral

neuropathies cause by physical injury or disease state, physical damage to the brain, physical damage to the spinal cord, stroke associated with brain damage, and neurological disorders relating to neurodegeneration.

5

18. The method of claim 16, wherein the neurological disorder is selected from the group consisting of Alzheimer's Disease, Parkinson's Disease, amyotrophic lateral sclerosis, and Huntington's Disease.

10

19. The method of claim 16, wherein the neurological disorder is Alzheimer's disease.

15

20. The method of claim 16, wherein the neurological disorder is Parkinson's disease.

21. The method of claim 16, wherein the neurological disorder is amyotrophic lateral sclerosis.

20

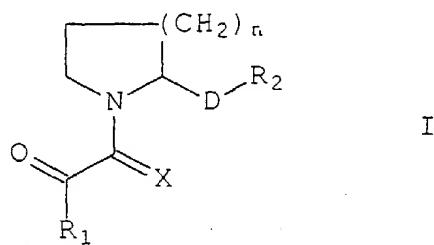
22. The method of claim 16, wherein the neurological disorder is Huntington's disease.

25

23. The method of claim 16, wherein the N-heterocyclic carboxylic acid or carboxylic acid isostere is non-immunosuppressive.

24. The method of claim 16, wherein the N-heterocyclic carboxylic acid or carboxylic acid isostere comprises a compound of formula (I):

30



where

n is 1-3;

X is either O or S;

R₁ is selected from the group consisting of C₁-C₉ straight or branched chain alkyl or alkenyl, C₂-C₉ straight or branched chain alkenyl, aryl, heteroaryl, carbocycle, or heterocycle;

D is a bond, or a C₁-C₁₀ straight or branched chain alkyl, C₂-C₁₀ alkenyl or C₂-C₁₀ alkynyl;

R₂ is carboxylic acid or a carboxylic acid isostere; and

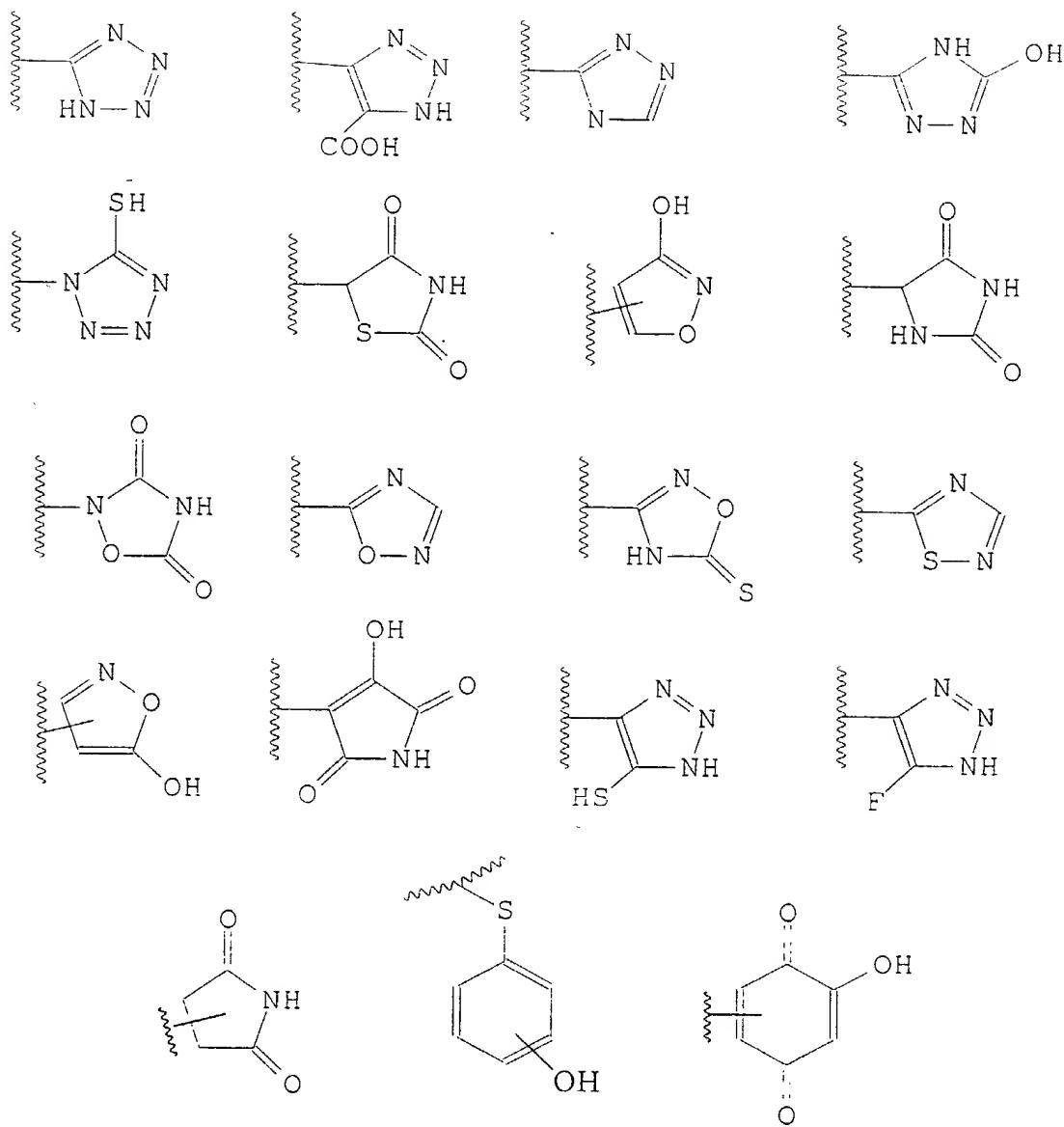
wherein said alkyl, alkenyl, alkynyl, aryl, heteroaryl, carbocycle, or heterocycle is optionally substituted with one or more substituents selected from R³, where

R³ is hydrogen, hydroxy, halo, haloalkyl, thiocarbonyl, alkoxy, alenoxy, alkylaryloxy, aryloxy, arylalkyloxy, cyano, nitro, imino, alkylamino, aminoalkyl, sulphydryl, thioalkyl, alkylthio, sulfonyl, C₁-C₆ straight or branched chain alkyl, C₂-C₆ straight or branched chain alkenyl or alkynyl, aryl, aralkyl, heteroaryl, carbocycle, heterocycle, and CO₂R⁷ where R⁷ is hydrogen or C₁-C₉ straight or branched chain alkyl or C₂-C₉ straight or branched chain alkenyl; or a pharmaceutically acceptable salt, ester, or solvate thereof.

25

25. The method of claim 24, wherein R₂ is a carbocycle or heterocycle containing any combination of CH₂, O, S, or N in any chemically stable oxidation state, wherein any of the atoms of said ring structure are optionally substituted in one or more positions with R³.

30
26. The method of claim 24, wherein R₂ is selected from the following group:



where the atoms of said ring structure may be optionally substituted at one or more positions with R³.

27. The method of claim 24, wherein R₂ is selected from the
5 group consisting of:

-COOH; -SO₃H; -SO₂HNR³; -PO₂(R³)₂; -CN; -PO₃(R³)₂; -OR³; -SR³;
-NHCOR³; -N(R³)₂; -CON(R³)₂; -CONH(O)R³; -CONHNHSO₂R³;
-COHNSO₂R³; and -CONR³CN..

10 28. The method of claim 16, wherein the N-heterocyclic carboxylic acid or carboxylic acid isostere compound is selected from the group consisting of compounds 1-139.

15 29. The method of claim 16, further comprising administering a neurotrophic factor different from formula (I).

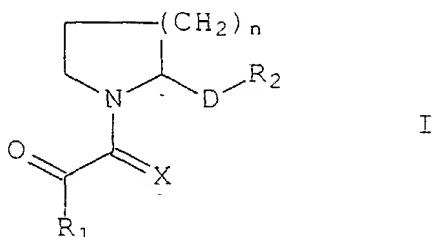
20 30. The method of claim 29, wherein said neurotrophic factor different from formula (I) is selected from the group consisting of neurotrophic growth factor, brain derived growth factor, glial derived growth factor, ciliary neurotrophic factor, insulin growth factor and active truncated derivatives thereof, acidic fibroblast growth factor, basic fibroblast growth factor, platelet-derived growth factors, neurotropin-3, and neurotropin 4/5.

25 31. A method of stimulating growth of damaged peripheral nerves, comprising:

30 administering to damaged peripheral nerves an effective amount of an N-heterocyclic carboxylic acid or carboxylic acid isostere to stimulate or promote growth of the damaged peripheral nerves.

35 32. The method of claim 31, wherein the N-heterocyclic carboxylic acid or carboxylic acid isostere is non-immunosuppressive.

33. The method of claim 31, wherein the N-heterocyclic carboxylic acid or carboxylic acid isostere comprises a compound of formula (I):



5 where

n is 1-3;

X is either O or S;

10 R₁ is selected from the group consisting of C₁-C₉, straight or branched chain alkyl or alkenyl, C₂-C₉, straight or branched chain alkenyl, aryl, heteroaryl, carbocycle, or heterocycle;

D is a bond, or a C₁-C₁₀ straight or branched chain alkyl, C₂-C₁₀ alkenyl or C₂-C₁₀ alkynyl;

R₂ is carboxylic acid or a carboxylic acid isostere;

15 and

wherein said alkyl, alkenyl, alkynyl, aryl, heteroaryl, carbocycle, or heterocycle is optionally substituted with one or more substituents selected from R³, where

20 R³ is hydrogen, hydroxy, halo, haloalkyl, thiocarbonyl, alkoxy, alkenoxy, alkylaryloxy, aryloxy, arylalkyloxy, cyano, nitro, imino, alkylamino, aminoalkyl, sulphydryl, thioalkyl, alkylthio, sulfonyl, C₁-C₆ straight or branched chain alkyl, C₂-C₆ straight or branched chain alkenyl or alkynyl, aryl, aralkyl, heteroaryl, carbocycle, heterocycle, and CO₂R⁷ where

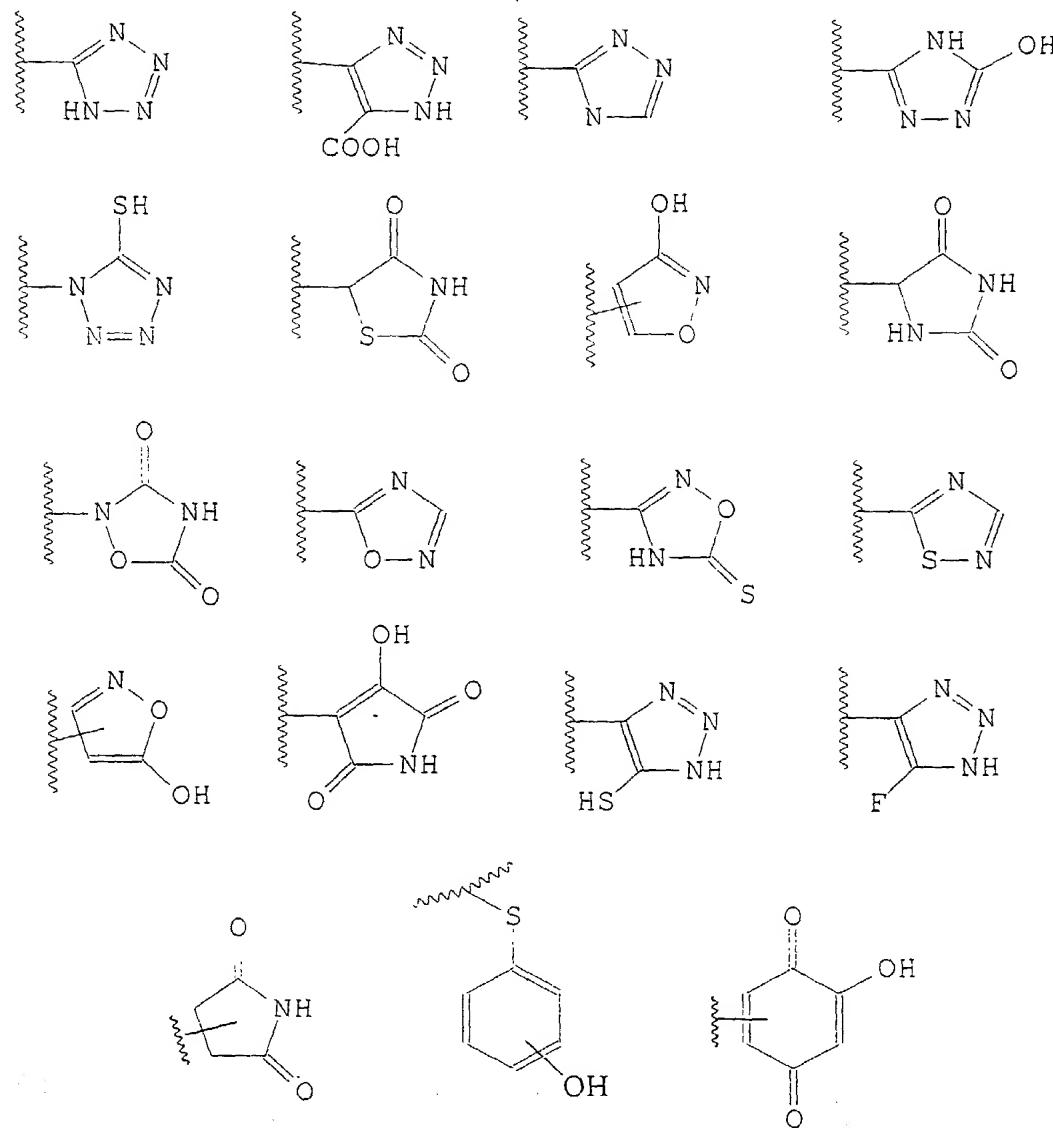
25 R⁷ is hydrogen or C₁-C₉, straight or branched chain alkyl or C₂-C₉, straight or branched chain alkenyl;

or a pharmaceutically acceptable salt, ester, or solvate

thereof.

34. The method of claim 33, wherein R₂ is a carbocycle or heterocycle containing any combination of CH₂, O, S, or N in any chemically stable oxidation state, wherein any of the atoms of said ring structure are optionally substituted in one or more positions with R³.

35. The method of claim 33, wherein R₂ is selected from the following group:



where the atoms of said ring structure may be optionally substituted at one or more positions with R³.

36. The method of claim 33, wherein R₂ is selected from the group consisting of:

-COOH; -SO₃H; -SO₂HNR³; -PO₂(R³)₂; -CN; -PO₃(R³)₂; -OR³; -SR³; -NHCOR³; -N(R³)₂; -CON(R³)₂; -CONH(O)R³; -CONHNHSO₂R³; -COHNSO₂R³; and -CONR³CN.

10 37. The method of claim 31, wherein the N-heterocyclic carboxylic acid or carboxylic acid isostere compound is selected from the group consisting of compounds 1-139.

15 38. The method of claim 31, further comprising administering a neurotrophic factor different from formula (I).

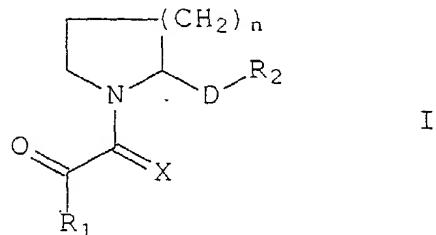
20 39. The method of claim 38, wherein said neurotrophic factor different from formula (I) is selected from the group consisting of neurotrophic growth factor, brain derived growth factor, glial derived growth factor, ciliary neurotrophic factor, insulin growth factor and active truncated derivatives thereof, acidic fibroblast growth factor, basic fibroblast growth factor, platelet-derived growth factors, neurotropin-3, and neurotropin 4/5.

25 40. A method for promoting neuronal regeneration and growth in animals, comprising:

30 administering to an animal an effective amount of a N-heterocyclic carboxylic acid or carboxylic acid isostere to promote neuronal regeneration.

35 41. The method of claim 40, wherein the N-heterocyclic carboxylic acid or carboxylic acid isostere is non-immunosuppressive.

42. The method of claim 40, wherein the N-heterocyclic carboxylic acid or carboxylic acid isostere comprises a compound of formula (I):



5 where

n is 1-3;

X is either O or S;

R₁ is selected from the group consisting of C₁-C₉, straight or branched chain alkyl or alkenyl, C₂-C₉, straight or branched chain alkenyl, aryl, heteroaryl, carbocycle, or heterocycle;

D is a bond, or a C₁-C₁₀ straight or branched chain alkyl, C₂-C₁₀ alkenyl or C₂-C₁₀ alkynyl;

R₂ is carboxylic acid or a carboxylic acid isostere;

15 and

wherein said alkyl, alkenyl, alkynyl, aryl, heteroaryl, carbocycle, or heterocycle is optionally substituted with one or more substituents selected from R³, where

R³ is hydrogen, hydroxy, halo, haloalkyl, thiocarbonyl, alkoxy, alkenoxy, alkylaryloxy, aryloxy, arylalkyloxy, cyano, nitro, imino, alkylamino, aminoalkyl, sulphydryl, thioalkyl, alkylthio, sulfonyl, C₁-C₆ straight or branched chain alkyl, C₂-C₆ straight or branched chain alkenyl or alkynyl, aryl, aralkyl, heteroaryl, carbocycle, heterocycle, and CO₂R⁷ where

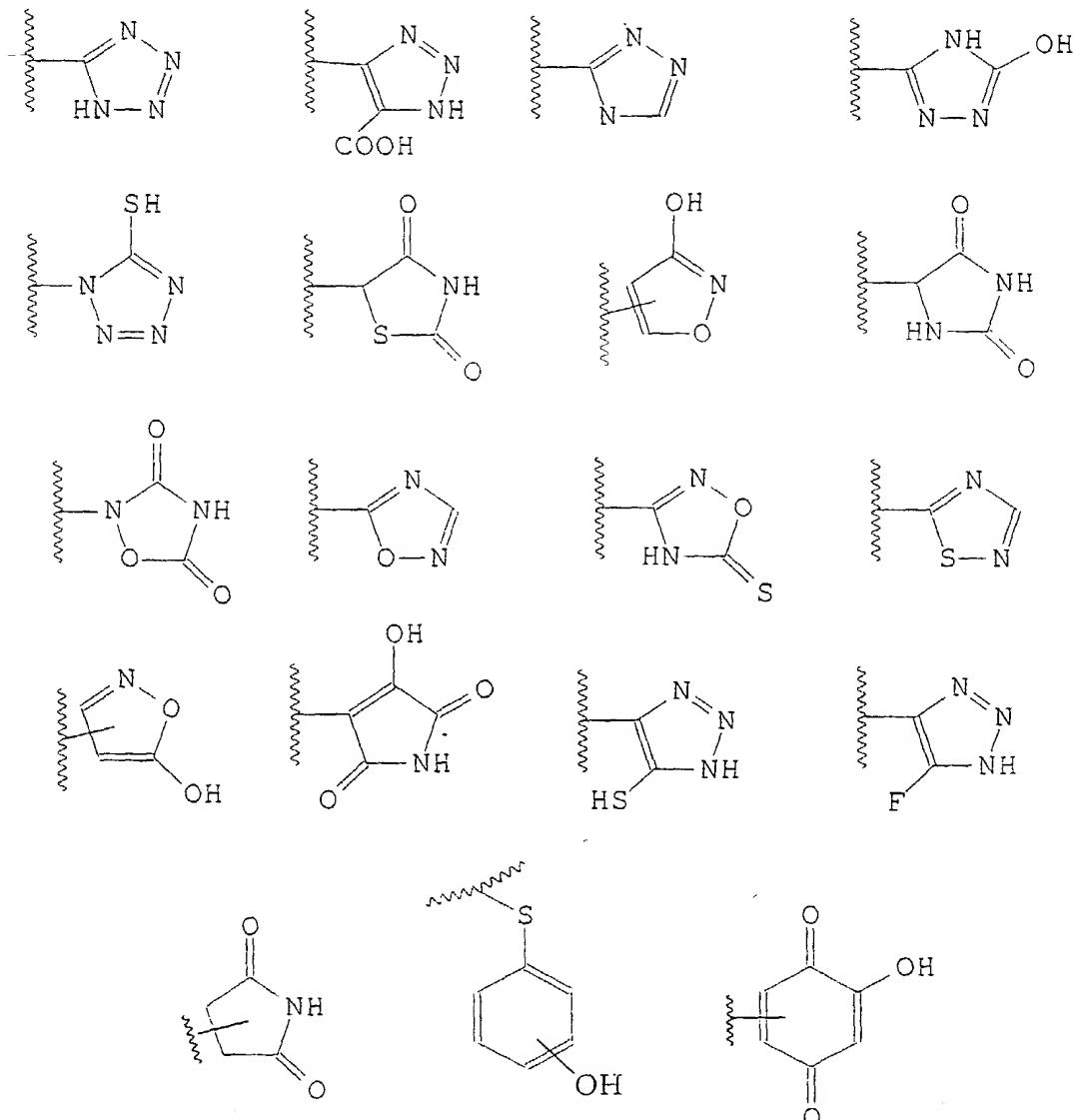
25 R⁷ is hydrogen or C₁-C₉, straight or branched chain alkyl or C₂-C₉, straight or branched chain alkenyl;

or a pharmaceutically acceptable salt, ester, or solvate

thereof.

43. The method of claim 42, wherein R_2 is a carbocycle or heterocycle containing any combination of CH_2 , O, S, or N in any chemically stable oxidation state, wherein any of the atoms of said ring structure are optionally substituted in one or more positions with R^3 .

44. The method of claim 42, wherein R_2 is selected from the following group:



where the atoms of said ring structure may be optionally substituted at one or more positions with R³.

45. The method of claim 42, wherein R₂ is selected from the group consisting of:

-COOH; -SO₃H; -SO₂HNR³; -PO₂(R³)₂; -CN; -PO₃(R³)₂; -OR³; -SR³; -NHCOR³; -N(R³)₂; -CON(R³)₂; -CONH(O)R³; -CONHNHSO₂R³; -COHNSO₂R³; and -CONR³CN..

10 46. The method of claim 40, wherein the N-heterocyclic carboxylic acid or carboxylic acid isostere compound is selected from the group consisting of compounds 1-139.

15 47. The method of claim 40, further comprising administering a neurotrophic factor different from formula (I).

20 48. The method of claim 47, wherein said neurotrophic factor different from formula (I) is selected from the group consisting of neurotrophic growth factor, brain derived growth factor, glial derived growth factor, ciliary neurotrophic factor, insulin growth factor and active truncated derivatives thereof, acidic fibroblast growth factor, basic fibroblast growth factor, platelet-derived growth factors, neurotropin-3, and neurotropin 4/5.

25

49. A method for preventing neurodegeneration in an animal, comprising:

30 administering to an animal an effective amount of a N-heterocyclic carboxylic acid or carboxylic acid isostere to prevent neurodegeneration.

50. The method of claim 49, wherein the neurodegeneration is Alzheimer's disease.

35 51. The method of claim 49, wherein the neurodegeneration

is Parkinson's disease.

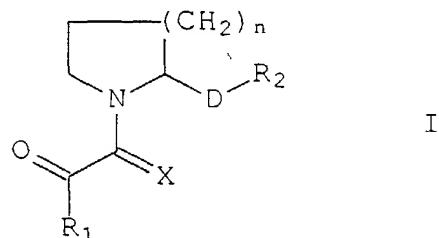
52. The method of claim 49, wherein the neurodegeneration is amyotrophic lateral sclerosis.

5

53. The method of claim 49, wherein the neurodegeneration is Huntington's Disease.

10 54. The method of claim 49, wherein the N-heterocyclic carboxylic acid or carboxylic acid isostere is non-immunosuppressive.

15 55. The method of claim 49, wherein the N-heterocyclic carboxylic acid or carboxylic acid isostere comprises a compound of formula (I):



where

n is 1-3;

X is either O or S;

20 R₁ is selected from the group consisting of C₁-C₉, straight or branched chain alkyl or alkenyl, C₂-C₉, straight or branched chain alkenyl, aryl, heteroaryl, carbocycle, or heterocycle;

D is a bond, or a C₁-C₁₀ straight or branched chain alkyl, 25 C₂-C₁₀ alkenyl or C₂-C₁₀ alkynyl;

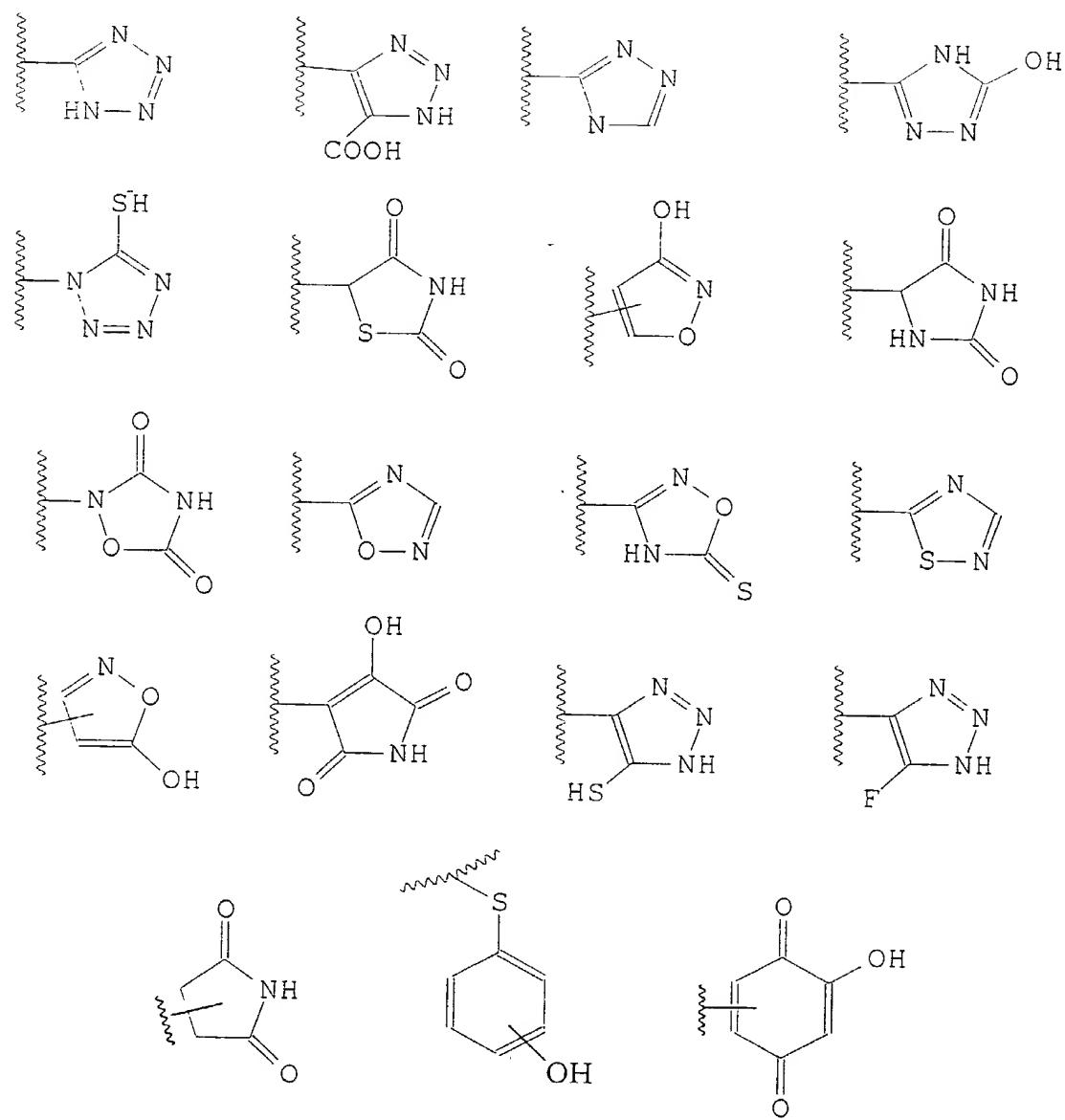
R₂ is carboxylic acid or a carboxylic acid isostere; and

wherein said alkyl, alkenyl, alkynyl, aryl, heteroaryl, carbocycle, or heterocycle is optionally substituted with one or more substituents selected from R³, where

R³ is hydrogen, hydroxy, halo, haloalkyl, thiocarbonyl, 5 alkoxy, alkenoxy, alkylaryloxy, aryloxy, arylalkyloxy, cyano, nitro, imino, alkylamino, aminoalkyl, sulfhydryl, thioalkyl, alkylthio, sulfonyl, C₁-C₆ straight or branched chain alkyl, C₂-C₆ straight or branched chain alkenyl or alkynyl, aryl, aralkyl, heteroaryl, carbocycle, heterocycle, and CO₂R⁷ where
10 R⁷ is hydrogen or C₁-C₉ straight or branched chain alkyl or C₂-C₉ straight or branched chain alkenyl;
or a pharmaceutically acceptable salt, ester, or solvate thereof.

15 56. The method of claim 55, wherein R₂ is a carbocycle or heterocycle containing any combination of CH₂, O, S, or N in any chemically stable oxidation state, wherein any of the atoms of said ring structure are optionally substituted in one or more positions with R³.

20 57. The method of claim 55, wherein R₂ is selected from the following group:



where the atoms of said ring structure may be optionally substituted at one or more positions with R³.

58. The method of claim 55, wherein R₂ is selected from the group consisting of:

-COOH; -SO₃H; -SO₂HNR³; -PO₂(R³)₂; -CN; -PO₃(R³)₂; -OR³; -SR³; -NHCOR³; -N(R³)₂; -CON(R³)₂; -CONH(O)R³; -CONHNHSO₂R³; -COHNSO₂R³; and -CONR³CN.

10 59. The method of claim 49, wherein the N-heterocyclic carboxylic acid or carboxylic acid isostere compound is
-- selected from the group consisting of compounds 1-139.

15 60. The method of claim 49, further comprising administering a neurotrophic factor different from formula (I).

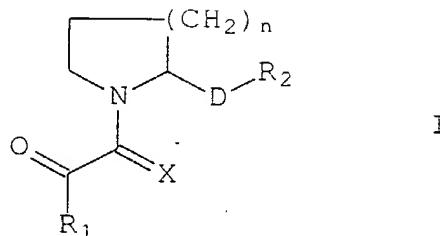
20 61. The method of claim 60, wherein said neurotrophic factor different from formula (I) is selected from the group consisting of neurotrophic growth factor, brain derived growth factor, glial derived growth factor, ciliary neurotrophic factor, insulin growth factor and active truncated derivatives thereof, acidic fibroblast growth factor, basic fibroblast growth factor, platelet-derived growth factors, neurotropin-3, and neurotropin 4/5.

25 62. A method for treating alopecia or promoting hair growth in an animal, which comprises administering to said animal an effective amount of an N-heterocyclic carboxylic acid or carboxylic acid isostere.

30 63. The method of claim 62, wherein the N-heterocyclic carboxylic acid or carboxylic acid isostere is non-immunosuppressive.

35 64. The method of claim 62, wherein the N-heterocyclic

carboxylic acid or carboxylic acid isostere is a compound of formula (I):



-- where

5 n is 1-3;

X is either O or S;

R₁ is selected from the group consisting of C₁-C₉, straight or branched chain alkyl or alkenyl, C₂-C₉, straight or branched chain alkenyl, aryl, heteroaryl, carbocycle, or heterocycle;

10 D is a bond, or a C₁-C₁₀ straight or branched chain alkyl, C₂-C₁₀ alkenyl or C₂-C₁₀ alkynyl;

R₂ is carboxylic acid or a carboxylic acid isostere; and

15 wherein said alkyl, alkenyl, alkynyl, aryl, heteroaryl, carbocycle, heterocycle, or carboxylic acid isostere is optionally substituted with one or more substituents selected from R³, where

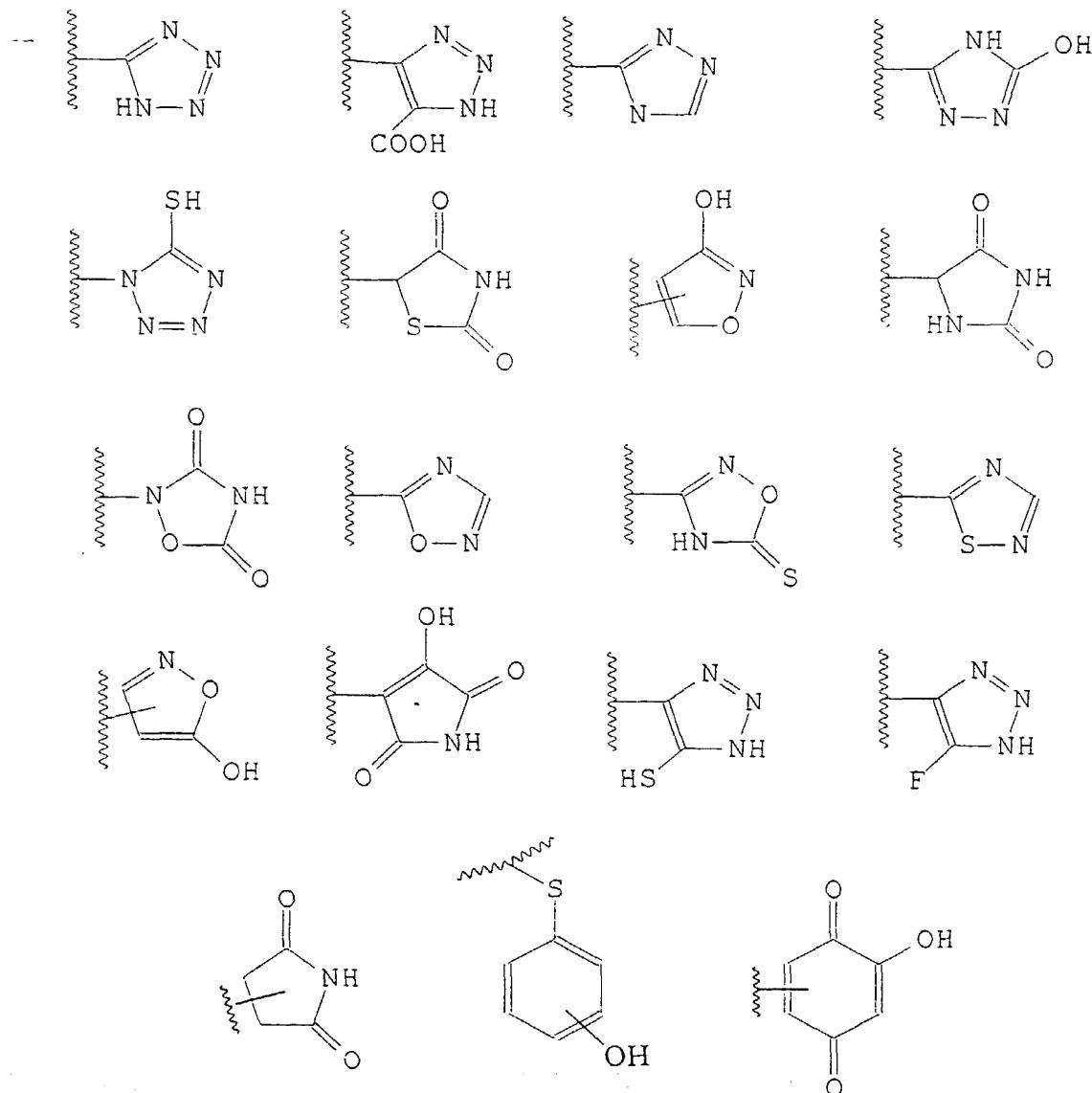
20 R³ is hydrogen, hydroxy, halo, haloalkyl, thiocarbonyl, alkoxy, alkenoxy, alkylaryloxy, aryloxy, arylalkyloxy, cyano, nitro, imino, alkylamino, aminoalkyl, sulphydryl, thioalkyl, alkylthio, sulfonyl, C₁-C₆ straight or branched chain alkyl, C₂-C₆ straight or branched chain alkenyl or alkynyl, aryl, aralkyl, heteroaryl, carbocycle, heterocycle, and CO₂R⁷ where

25 R⁷ is hydrogen or C₁-C₉, straight or branched chain alkyl or C₂-C₉, straight or branched chain alkenyl; or a pharmaceutically acceptable salt, ester, or solvate

thereof.

65. The method of claim 64, wherein R₂ is a carbocycle or heterocycle containing any combination of CH₂, O, S, or N in any chemically stable oxidation state, wherein any of the atoms of said ring structure are optionally substituted in one or more positions with R³.

66. The method of claim 64, wherein R₂ is selected from the following group:



where the atoms of said ring structure may be optionally substituted at one or more positions with R³.

67. The method of claim 64, wherein R₂ is selected from the group consisting of

-COOH; -SO₃H; -SO₂HNR³; -PO₂(R³)₂; -CN; -PO₃(R³)₂; -OR³; -SR³; -NHCOR³; -N(R³)₂; -CON(R³)₂; -CONH(O)R³; -CONHNHSO₂R³; -COHNSO₂R³; and -CONR³CN.

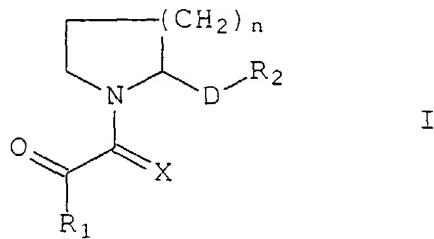
68. The method of claim 62, wherein the carboxylic acid or carboxylic acid isostere is selected from the group consisting of compounds 1-139.

69. A pharmaceutical composition comprising:

(i) an effective amount of a N-heterocyclic carboxylic acid or carboxylic acid isostere for treating alopecia or promoting hair growth in an animal; and
(ii) a pharmaceutically acceptable carrier.

70. The pharmaceutical composition of claim 69, wherein the N-heterocyclic carboxylic acid or carboxylic acid isostere is non-immunosuppressive.

71. The composition of claim 69, wherein the carboxylic acid or carboxylic acid isostere is a compound of formula (I):



where

n is 1-3;

X is either O or S;

5 R₁ is selected from the group consisting of C₁-C₉, straight or branched chain alkyl or alkenyl, C₂-C₉, straight or branched chain alkenyl, aryl, heteroaryl, carbocycle, or heterocycle;

D is a bond, or a C₁-C₁₀ straight or branched chain alkyl, C₂-C₁₀ alkenyl or C₂-C₁₀ alkynyl;

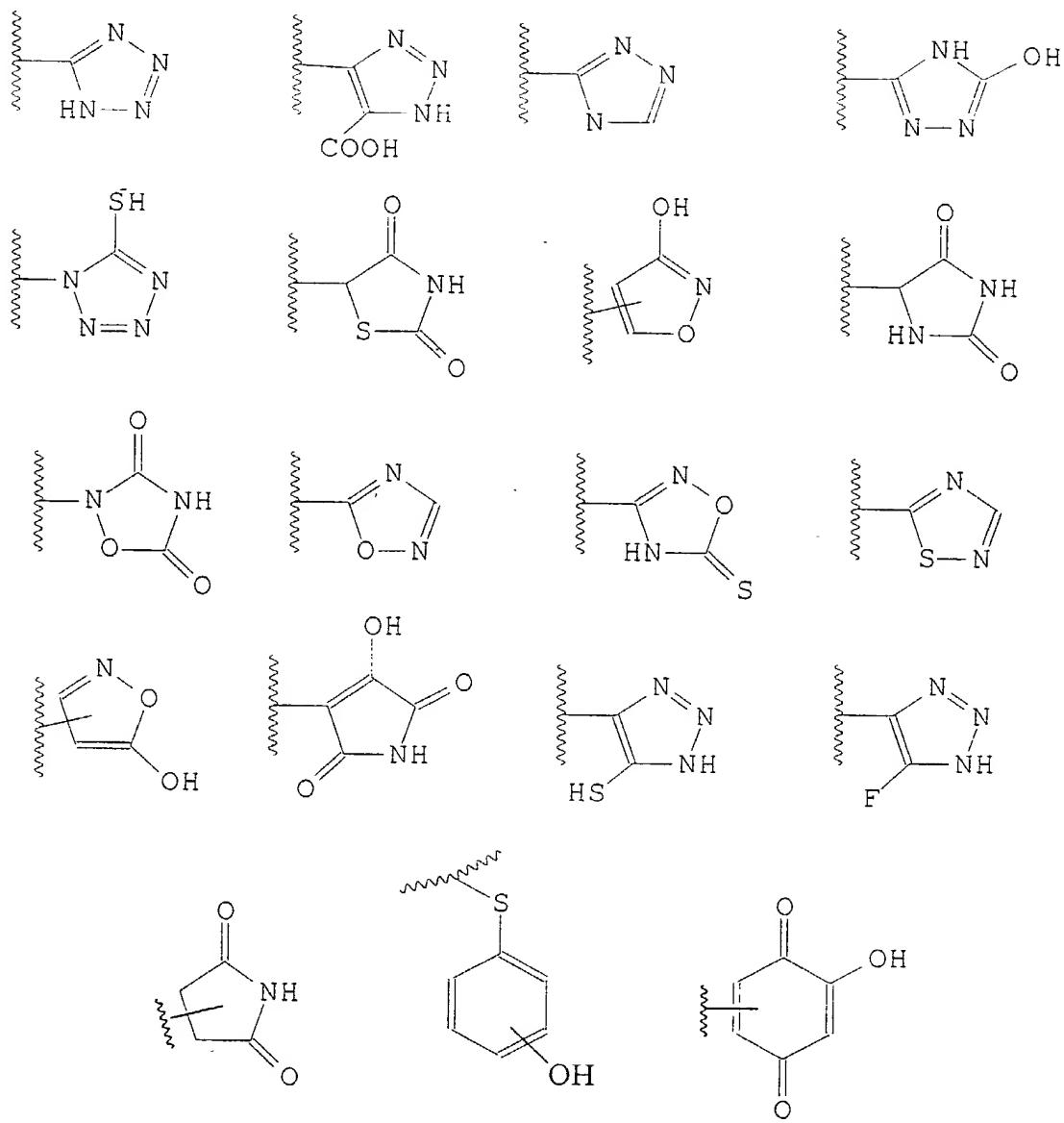
10 R₂ is carboxylic acid or a carboxylic acid isostere; and

wherein said alkyl, alkenyl, alkynyl, aryl, heteroaryl, carbocycle, heterocycle, or carboxylic acid isostere is optionally substituted with one or more substituents selected 15 from R³, where

R³ is hydrogen, hydroxy, halo, haloalkyl, thiocarbonyl, alkoxy, alenoxy, alkylaryloxy, aryloxy, arylalkyloxy, cyano, nitro, imino, alkylamino, aminoalkyl, sulfhydryl, thioalkyl, alkylthio, sulfonyl, C₁-C₆ straight or branched chain alkyl, 20 C₂-C₆ straight or branched chain alkenyl or alkynyl, aryl, aralkyl, heteroaryl, carbocycle, heterocycle, and CO₂R⁷ where R⁷ is hydrogen or C₁-C₉ straight or branched chain alkyl or C₂-C₉ straight or branched chain alkenyl; or a pharmaceutically acceptable salt, ester, or solvate 25 thereof.

72. The composition of claim 71, wherein R₂ is a carbocycle or heterocycle containing any combination of CH₂, O, S, or N in any chemically stable oxidation state, wherein any of the 30 atoms of said ring structure are optionally substituted in one or more positions with R³.

73. The composition of claim 71, wherein R₂ is selected from the following group:



where the atoms of said ring structure may be optionally substituted at one or more positions with R³.

74. The composition of claim 71, wherein R₂ is selected from
5 the group consisting of:

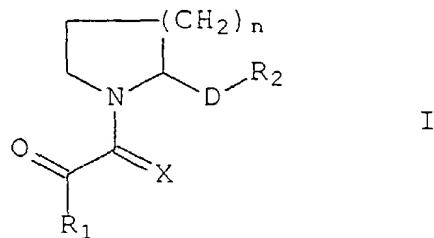
-COOH; -SO₃H; -SO₂HNR³; -PO₂(R³)₂; -CN; -PO₃(R³)₂; -OR³; -SR³;
-NHCOR³; -N(R³)₂; -CON(R³)₂; -CONH(O)R³; -CONHNHSO₂R³;
-COHNSO₂R³; and -CONR³CN.

10 75. The composition of claim 69, wherein the carboxylic acid
or carboxylic acid isostere is selected from the group
consisting of compounds 1-139.

15 76. A method for treating a vision disorder, improving
vision, treating memory impairment, or enhancing memory
performance in an animal, which comprises administering to
said animal an effective amount of an N-heterocyclic
carboxylic acid or carboxylic acid isostere.

20 77. The method of claim 76, wherein the N-heterocyclic
carboxylic acid or carboxylic acid isostere is non-
immunosuppressive.

25 78. The method of claim 76, wherein the N-heterocyclic
carboxylic acid or carboxylic acid isostere is a compound of
formula (I):



where

n is 1-3;

X is either O or S;

R₁ is selected from the group consisting of C₁-C₉ straight or branched chain alkyl or alkenyl, C₂-C₉ straight or branched chain alkenyl, aryl, heteroaryl, carbocycle, or heterocycle;

D is a bond, or a C₁-C₁₀ straight or branched chain alkyl, C₂-C₁₀ alkenyl or C₂-C₁₀ alkynyl;

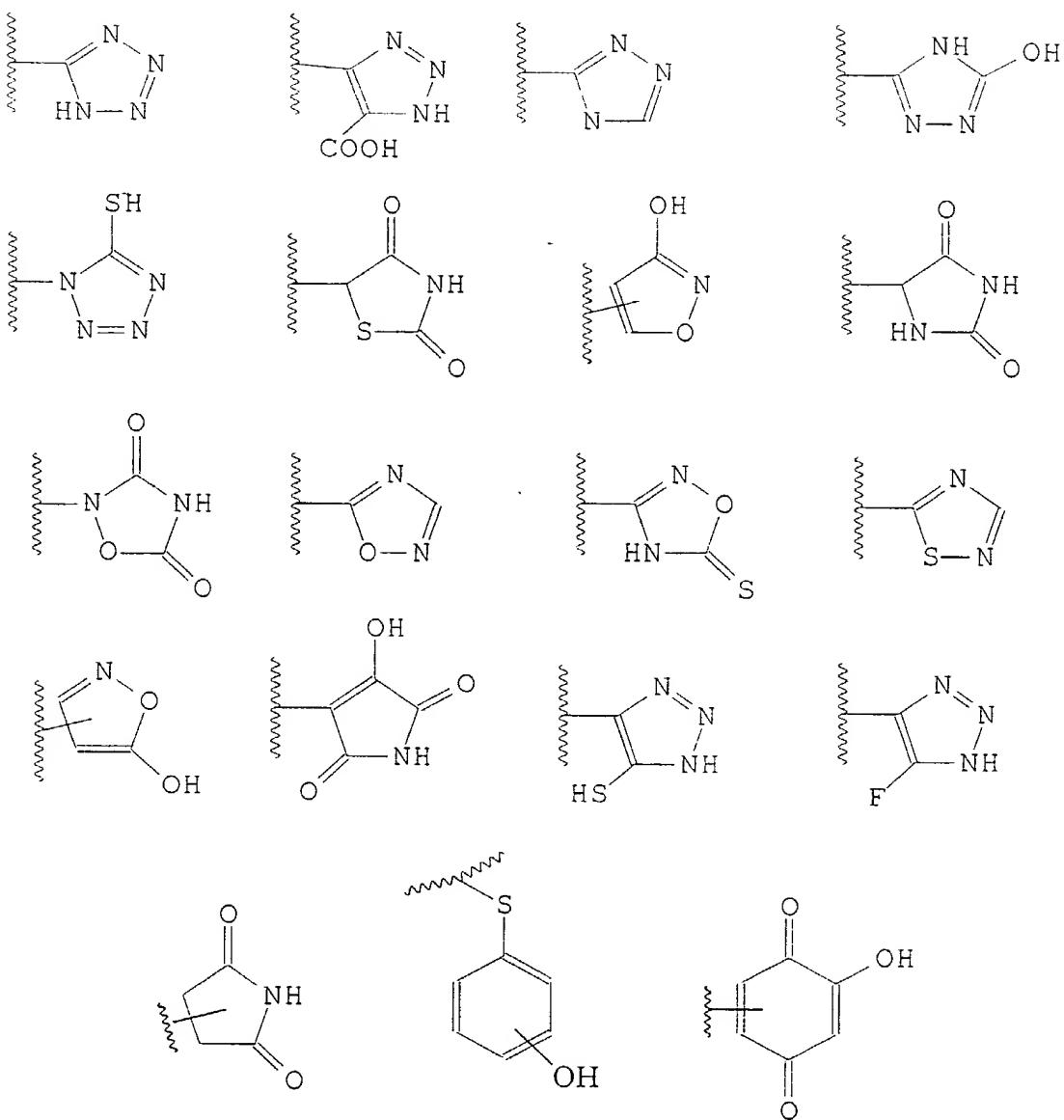
R₂ is carboxylic acid or a carboxylic acid isostere; and

wherein said alkyl, alkenyl, alkynyl, aryl, heteroaryl, carbocycle, heterocycle, or carboxylic acid isostere is optionally substituted with one or more substituents selected from R³, where

R³ is hydrogen, hydroxy, halo, haloalkyl, thiocarbonyl, alkoxy, alkenoxy, alkylaryloxy, aryloxy, arylalkyloxy, cyano, nitro, imino, alkylamino, aminoalkyl, sulfhydryl, thioalkyl, alkylthio, sulfonyl, C₁-C₆ straight or branched chain alkyl, C₂-C₆ straight or branched chain alkenyl or alkynyl, aryl, aralkyl, heteroaryl, carbocycle, heterocycle, and CO₂R⁷ where R⁷ is hydrogen or C₁-C₉ straight or branched chain alkyl or C₂-C₉ straight or branched chain alkenyl; or a pharmaceutically acceptable salt, ester, or solvate thereof.

79. The method of claim 78, wherein R₂ is a carbocycle or heterocycle containing any combination of CH₂, O, S, or N in any chemically stable oxidation state, wherein any of the atoms of said ring structure are optionally substituted in one or more positions with R³.

80. The method of claim 78, wherein R₂ is selected from the following group:



where the atoms of said ring structure may be optionally substituted at one or more positions with R³.

81. The method of claim 78, wherein R₂ is selected from the
5 group consisting of

-COOH; -SO₃H; -SO₂HNR³; -PO₂(R³)₂; -CN; -PO₃(R³)₂; -OR³; -SR³;
-NHCOR³; -N(R³)₂; -CON(R³)₂; -CONH(O)R³; -CONHNHSO₂R³;
-COHNSO₂R³; and -CONR³CN.

10 82. The method of claim 76, wherein the carboxylic acid or carboxylic acid isostere is selected from the group consisting of compounds 1-139.